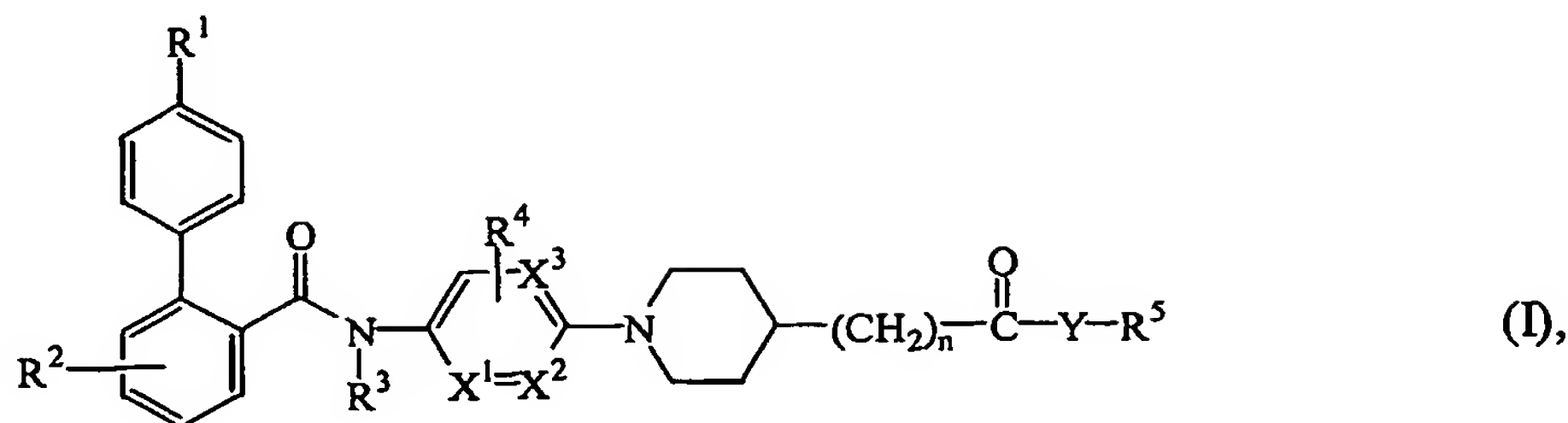


Claims

1. A compound of formula (I)



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the *N*-oxides, the pharmaceutically acceptable acid addition salts and the stereochemically isomeric forms thereof, wherein

$R^1$  is hydrogen,  $C_{1-4}$ alkyl, halo, or polyhalo $C_{1-4}$ alkyl;

$R^2$  is hydrogen,  $C_{1-4}$ alkyl, halo, or polyhalo $C_{1-4}$ alkyl;

10  $R^3$  is hydrogen or  $C_{1-4}$ alkyl;

$R^4$  is hydrogen,  $C_{1-4}$ alkyl, or halo;

$n$  is an integer 0, or 1;

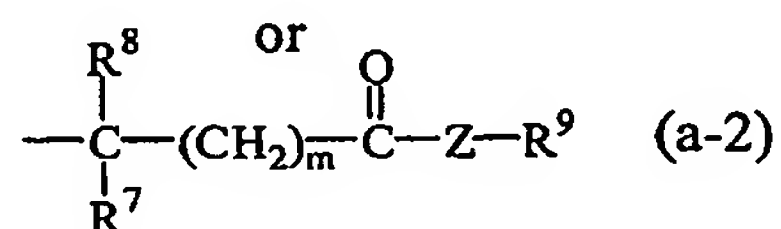
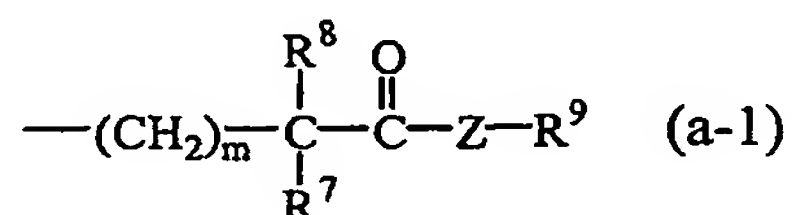
$X^1$  is carbon and  $X^2$  is carbon; or  $X^1$  is nitrogen and  $X^2$  is carbon;

or  $X^1$  is carbon and  $X^2$  is nitrogen;

15  $X^3$  is carbon or nitrogen;

$Y$  represents O, or  $NR^6$  wherein  $R^6$  is hydrogen or  $C_{1-4}$ alkyl;

$R^5$  represents a radical of formula



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wherein

$m$  is an integer 0, 1, or 2;

$Z$  is O or NH;

$R^7$  is hydrogen,

$C_{1-6}$ alkyl;

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$C_{1-6}$ alkyl substituted with hydroxy, amino, mono- or

di( $C_{1-4}$ alkyl)amino,  $C_{1-4}$ alkyloxycarbonyl, aminocarbonyl, aryl or heteroaryl;

$C_{1-4}$ alkyl-O- $C_{1-4}$ alkyl;

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C<sub>1-4</sub>alkyl-S-C<sub>1-4</sub>alkyl; or  
aryl;

R<sup>8</sup> is hydrogen or C<sub>1-6</sub>alkyl;

R<sup>9</sup> is hydrogen, C<sub>1-4</sub>alkyl, aryl<sup>1</sup>, or C<sub>1-4</sub>alkyl substituted with aryl<sup>1</sup>;

5 or when Y represents NR<sup>6</sup> the radicals R<sup>5</sup> and R<sup>6</sup> may be taken together with the nitrogen to which they are attached to form pyrrolidinyl substituted with C<sub>1-4</sub>alkyloxycarbonyl and optionally further substituted with hydroxy; or piperidinyl substituted with C<sub>1-4</sub>alkyloxycarbonyl;

10 aryl is phenyl; phenyl substituted with one, two or three substituents each independently selected from C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxy, halo, hydroxy, nitro, cyano, C<sub>1-4</sub>alkyloxycarbonyl, trifluoromethyl, or trifluoromethoxy; or benzo[1,3]dioxolyl;

15 aryl<sup>1</sup> is phenyl; phenyl substituted with one, two or three substituents each independently selected from C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxy, halo, hydroxy, nitro, cyano, C<sub>1-4</sub>alkyloxycarbonyl, trifluoromethyl, or trifluoromethoxy; and heteroaryl is imidazolyl, thiazolyl, indolyl, or pyridinyl.

2. A compound as claimed in claim 1 wherein X<sup>1</sup>, X<sup>2</sup> and X<sup>3</sup> are carbon.

20 3. A compound as claimed in claim 1 wherein R<sup>1</sup> is trifluoromethyl; R<sup>2</sup> is hydrogen; R<sup>3</sup> is hydrogen; R<sup>4</sup> is hydrogen; X<sup>1</sup>, X<sup>2</sup> and X<sup>3</sup> are carbon; n is the integer 1; Y represents NR<sup>6</sup> wherein R<sup>6</sup> is hydrogen or methyl; and R<sup>5</sup> is a radical of formula (a-1) wherein m is the integer 0.

25 4. A compound as claimed in claim 1 wherein R<sup>1</sup> is trifluoromethyl; R<sup>2</sup> is hydrogen; R<sup>3</sup> is hydrogen; R<sup>4</sup> is hydrogen; X<sup>1</sup>, X<sup>2</sup> and X<sup>3</sup> are carbon; n is the integer 1; Y represents NR<sup>6</sup> wherein R<sup>6</sup> is hydrogen or methyl; and R<sup>5</sup> is a radical of formula (a-1) wherein m is the integer 1.

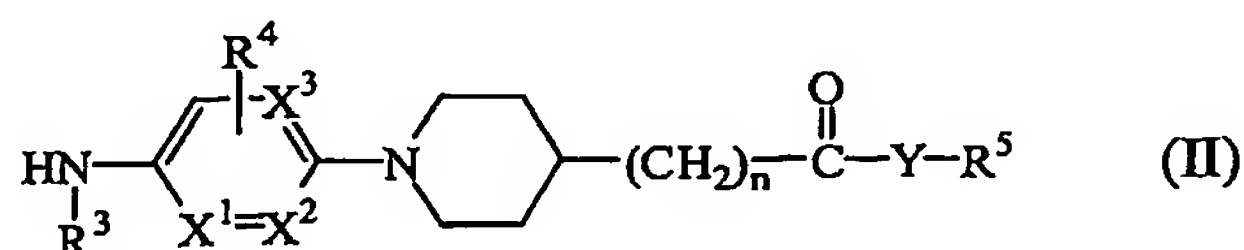
30 5. A compound as claimed in claim 1 wherein R<sup>1</sup> is trifluoromethyl; R<sup>2</sup> is hydrogen; R<sup>3</sup> is hydrogen; R<sup>4</sup> is hydrogen; X<sup>1</sup>, X<sup>2</sup> and X<sup>3</sup> are carbon; n is the integer 1; Y represents NR<sup>6</sup> wherein R<sup>6</sup> is hydrogen or methyl; and R<sup>5</sup> is a radical of formula (a-2) wherein m is the integer 1.

35 6. A compound as claimed in claim 1 wherein R<sup>1</sup> is trifluoromethyl; R<sup>2</sup> is hydrogen; R<sup>3</sup> is hydrogen; R<sup>4</sup> is hydrogen; X<sup>1</sup>, X<sup>2</sup> and X<sup>3</sup> are carbon; n is the integer 1; Y represents NR<sup>6</sup> and R<sup>5</sup> and R<sup>6</sup> are taken together with the nitrogen to which they are attached to form pyrrolidinyl substituted with C<sub>1-4</sub>alkyloxycarbonyl and optionally

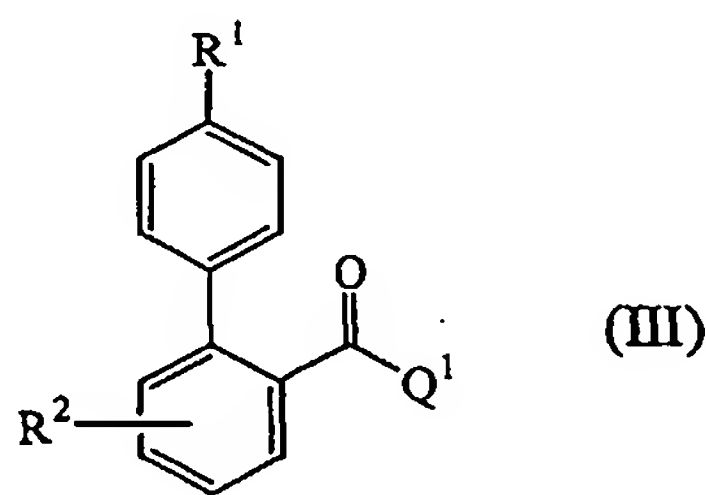
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further substituted with hydroxy, or piperidinyl substituted with C<sub>1-4</sub>alkyloxy-carbonyl.

7. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically active amount of a compound as claimed in any of claims 1 to 6.
8. A process for preparing a pharmaceutical composition as claimed in claim 7 wherein a therapeutically active amount of a compound as claimed in any of claims 1 to 6 is intimately mixed with a pharmaceutically acceptable carrier.
9. A compound as claimed in any of claims 1 to 6 for use as a medicine.
10. A process for preparing a compound of formula (I) wherein
  - a) an intermediate of formula (II), wherein R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup>, Y, n, X<sup>1</sup>, X<sup>2</sup> and X<sup>3</sup> are defined as in claim 1,



is reacted with a biphenylcarboxylic acid or halide having the formula (III), wherein R<sup>1</sup> and R<sup>2</sup> are as defined in formula (I) and Q<sup>1</sup> is selected from hydroxy and halo, in at least one reaction-inert solvent and optionally in the presence of a suitable base



- b) or, compounds of formula (I) are converted into each other following art-known transformation reactions; or if desired; a compound of formula (I) is converted into an acid addition salt, or conversely, an acid addition salt of a compound of formula (I) is converted into a free base form with alkali; and, if desired, preparing stereochemically isomeric forms thereof.